

#### (1) GENERAL INFORMATION:

- (i) APPLICANT: TSYRLOVA, IFENA WOLPE, STEPHEN D.
- (ii) TITLE OF INVENTION: INHIBITOR OF STEM CELL PROLIFERATION AND USES THEREOF
- (iii) NUMBER OF SEQUENCES: 37
- (iv) CORRESPONDENCE ADDRESS:
  - (A) ADDRESSEE: NIMON & VANDERHYE P.C.
  - (B) STREET: 1100 FORTH GLERE ROAD
  - (C) CITY: ARLINGTON
  - (D) STATE: UIRGINIA
  - (E) COUNTRY: U.S.A.
  - (F) DIF: 20001-4704
  - (v) COMPUTER READABLE FORM:
    - (A) MEDIUM TYPE: Ploppy disk
    - (B) COMPUTER: IBM PC compatible
    - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
    - (D) SOFTWARE: PatentIr. Release #1.0, Version #1.30
- (v1) CURRENT APPLACATION DATA:
  - (A) APPLICATION NUMBER: US 08:832,443
  - (B) FILING DATE: UN-APR-1997
  - (C) CLASSIFICATION:
- (viii) ATTORNEY AGENT INFORMATION:
  - (A) NAME: BYRNE, THOMAS E.
  - (B) REGISTRATION NUMBER: 32,205
  - (C) REFERENCE/DOCKET NUMBER: 1331-222
  - (ix) TELECOMMUNICATION INFORMATION:
    - (A) TELEPHONE: (70.) 816-4000
    - (B) TELEFAM: (103) 816-4100
  - TINA BRATTON BOARS NO ID NOR IX
    - AND SEQUENCE CHARACTERISTICS:
      - (A) LENGTH: 13 amino acids
      - (B) FYPE: amino acid
      - (C) STRANDEDNESS:
      - po ToPoloGY linear
    - er in the title over libe

- (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 15 amino acids (E) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (x1) SEQUENCE DESCRIPTION: SEQ ID NO:2: Cys Phe Pro His Phe Asp Leu Ser His Gly Ser Ala Gln Val Cys 10
- (2) INFORMATION FOR SEQ ID NO:3:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 19 amino acids
    - (E) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (11) MOLECULE TYPE: peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:
  - Asp Ala Leu Thr Asn Ala Val Ala His Val Asp Asp Met Pro Asn Ala 10

Leu Ser Ala

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- ACH INFORMATION FOR SEQ ID NO:4:
  - . REQUENCE CHARACTERISTICS:
    - A CLEDER BY DE Amino actifs on TYPE: amino actif one STPANDEDNESS:

    - (D) TOPOLOGY: linear
  - (i) MOLECULE TYPE: peptide

ing matter interest in the control of a

- (C) STRANDEDNESS:
- (D) TOPOLOGY, linear
- All: MOLECULE TYPE, peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

Leu Val Val Tyr Pro Trp Thr Gln Arg

- (2) INFORMATION FOR SEQ ID NO:6:
  - (i) SEQUENCE CHARACTER'STICS:
    - (A) LENGTH: 8 amine acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

Leu Val Val Tyr Pro Trp Thr Gln 1

- (2) INFORMATION FOR SEQ ID NO:7:
  - (i) SEQUENCE CHAFACTERISTICS:
    - (A) LENGTH: " amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - HILL MOLECULE TYPE: peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:

let Tal Val Tyr Pro Trp Thr
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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:8: Leu Val Val Tyr Pro Trp (2) INFORMATION FOR SEQ ID NO:9: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MODECULE TYPE, peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:9: Leu Val Val Tyr Pro

- (2) INFORMATION FOR SEQ ID NO:10:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 3 amino acids
    - (B) TYPE: amino acid
    - (0) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:10:
  - Tal Val Tyr Pro Trp The Glr
  - COR FMAILCO FOR SE, ID NOT LOS
    - (i) SEQUENCE CHARACTERISTICS:
      - (A) LENGTH: 7 amino acids
      - (B) TYPE: amino acid
      - (C) STRANDEDNESS:
      - (D) TOPOLOGY: linear
    - ii Moi Ford PortPE: peptide

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(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 6 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ 1D NO:12: Tyr Pro Trp Thr Gln Arg (2) INFORMATION FOR SEQ ID NO:13: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:13: Tyr Pro Trp Thr Gln (2) INFORMATION FOR SEQ ID NO:14: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 6 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: ch: TOPOLOGY: linear I MY ENTE TYPE bentile

(x:) SEQUENCE DESCRIPTION: SEQ ID NO:14:

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## (ii) MOLECULE TYPE: DNA (genomic)

	(xi) Si	EQUENCE DESC	CRIPTION: SE	EQ ID NO:15:	:		
<b>31.6</b> 0	TGICIC	CTGCCGACAA	GACCAACGTC	AAGGCCGCCT	GGGGTAAGGT	CGGCGCGCAC	юC
GCTC	GCGAGT	ATGGTGCGGA	GGCCCTGGAG	AGGATGTTCC	TGTCCTTCCC	CACCACCAAG	1: C
ACCI	ACTICC	CGCACTTCGA	CCTGAGCCAC	GGGTGTGCCC	AGGTTAAGGG	CCACGGCAAG	180
AAGG	TGGCCG	ACGCGCTGAC	CAACGCUGTG	GCGCACG PGG	ACGACATGCC	CAACGCGCTG	2.,0
TCCG	SCCCTCA	асаластасA	CGCGCACAAG	CTTCGGGTGG	ACCCGGTCAA	CITCAAGCTC	3.10
CTAP	GCCACT	GCCTGCTGGT	GACCCTGGCC	GOOGAGGTGG	CCGCCGAGTT	CACCCCTGCG	3 % 0
GTGC	CACGCCT	CCCTGGACAA	GTTCCTGGCT	TCTGTGAGCA	CCGTGCTGAC	CTCCAAATAC	41.0
CGT							423

## (2) INFORMATION FOR SEQ ID NO:16:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 141 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:

- Tal Lou Ser Pro Ala Asp Lys Thr Ash Val Lys Ala Ala Trp Gly Lys L
- The dry Ala His Ala dry dre Typ (try Ala Her Ala Lee dle Abd Met Dr
- The led Ser Phe Pro Thr Thr Lys Thr Tyr Phe Pro His Phe Asp Led 35 40 45
- Ser His Gly Ser Ala Gln Val Lys Gly His Gly Lys Lys Val Ala Asp
- Ala ton Thr Ash Ala Mal Ala His Mal Asp Asp Mor Pro Ash Ala Lou

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Leu Ala Ser Val Ser Thr Val Leu Thr Ser Lys Tyr Arg 130 135 140

### (2) INFORMATION FOR SEQ ID NO:17:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 438 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:17:

(	GTGCACCTGA	CTCCTGAGGA	GAAGTCTGCC	GTTACTGCCC	TGTGGGGCAA	GGTGAACGTG	ъ́0
(	GATGAAGTTG	GTGGTGAGGC	CCTGGGCAGG	CTGCTGGTGG	TCTACCTTTG	GACCCAGAGG	120
	PTCTTTGAGT	CCTTTGGGGGA	TOTGTCCACT	CCTGATGCTG	TTATGGGCAA	CCCTAAGGTG	130
-	AAGGCTCATG	GCAAGAAACT	GCTCCGTGCC	PTTAGTGATG	GCCTGGCTCA	CCTGGACAAC	240
ı	UTCAAGGGGA	COTTTGCCAC	ACTGAGTGAG	CTGCACTGT3	ACAAGCIGCA	CGTGGATCCT	300
{	GAGAACTTCA	GGCTGCTGGG	CAACGTGCTG	GTCTGTGTGC	TGGCCCATCA	CTTTGGCAAA	360
{	GAATTCACCC	CACCAGTGCA	GGCTGCCTAT	CAGAAAGTGG	TGGCTGGTGT	GGCTAATGCC	420
(	CTGGCCCACA	AGTATCAC					438

- (E) INFORMATION FOR SEQ ID NO:13:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 146 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) ToPOLOGY, linear
  - HE MOLECULE TYPE, peptide:
  - MEN SEQUENCE DESCRIPTION: SE, ID N :18:
  - Tal His ion The Broding in type Ber Ala Dal The Ala Ion Tip by

Lys Lys Val Leu Gly Ala Phe Ser Asp Gly Leu Ala His Leu Asp Asn 65 70 75 80

Leu Lys Gly Thr Phe Ala Thr Leu Ser Glu Leu His Cys Asp Lys Leu 85 90 95

His Val Asp Pro Glu Asn Phe Arg Leu Leu Gly Asn Val Leu Val Cys 100 105 110

Val Leu Ala His His Phe Gly Lys Glu Phe Thr Pro Pro Val Gln Ala 115 120 125

Ala Tyr Gln Lys Val Val Ala Gly Val Ala Asn Ala Leu Ala His Lys 130 135 140

Tyr His

### (2) INFORMATION FOR SEQ ID NO:19:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 141 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:19:
- Val Leu Ser Gly Glu Asp Lys Ser Asn Ile Lys Ala Ala Trp Gly Lys 1 5 10 15
- Ile Gly Gly His Gly Ala Glu Tyr Gly Ala Glu Ala Leu Glu Arg Met 25 30
- Pho Ala Ser she Pro Thr Inc Lys Inc Tyr Pho Fro His Pho Asp Web
- rvo Hir Bly Nor Ala Bln Mal tys Bly Hir Bly Tyr Lys Mal Ala Asy 50 55 60
- Ala Leu Ala Ser Ala Ala Gly His Leu Asp Asp Leu Pro Gly Ala Leu 65 70 75 80
- Sor Ala Len Sor Asp Leu His Ala His Lys Leu Arg Val Asp Pro Val

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- (2) INFORMATION FOR SEQ ID NO:20:
  - (i) SEQUENCE CHAPACTERISTICS:
    - (A) LENGTH: 146 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:20:
  - Val His Lei Thr Asp Ala Glu Lys Ala Ala Val Ser Cys Leu Trp Gly
    1 10 15
  - Lys Val Asn Ser Asp Glu Val Gly Gly Glu Ala Leu Gly Arg Leu Leu 20 25 30
  - Val Val Tyr Pro Trp Thr Gln Arg Tyr Phe Asp Ser Phe Gly Asp Leu 35 40 45
  - Ser Ser Ala Ser Ala Ile Met Gly Asr. Ala Lys Val Lys Ala His Gly 50 55 60
  - Lys Lys Val Ile Thr Ala Phe Asn Asp Gly Leu Asn His Leu Asp Ser 65 70 75 80
  - Leu Lys Gly Thr Phe Ala Ser Leu Ser Glu Leu His Cys Asp Lys Leu 85 90 95
  - His Val Asp Pro Glu Asn Phe Arg Leu Leu Gly Asn Met Ile Val Ile 100 105 110
  - Val Leu Gly His His Leu Gly Lys Asp Phe Thr Pro Ala Ala Gln Ala 115 120 125
  - Ala Phe G : Mys Val Val Ala Gly Val Ala Thr Ala Leu Ala His Lys 130 135 140
  - iyr His
- (1) INFORMATION FOR SEQ ID NO:21:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 141 amino acids
    - UB: TYPE amino acid
    - (67) STRANDEDNESS:

10

Val Gly Gly Gln Ala Gly Ala His Gly Ala Glu Ala Leu Glu Arg Met 20 25 30

Phe Lea Gly Phe Pro Thr Thr Lys Thr Tyr Phe Pro His Phe Ash Lea 35 40 45

Ser His Gly Ser Asp Gln Val Lys Ala His Gly Gln Lys Val Ala Asp 50 55 60

Ala Leu Thr Lys Ala Val Gly His Leu Asp Asp Leu Pro Gly Ala Leu 65 70 75 80

Ser Ala Leu Ser Asp Leu His Ala His Lys Leu Arg Val Asp Pro Val 85 90 95

His Pro Asp Asp Phe Asn Pro Ser Val His Ala Ser Leu Asp Lys Phe 115 120 125

Leu Ala Asn Val Ser Thr Val Leu Thr Ser Lys Tyr Arg 130 135 140

# (2) INFORMATION FOR SEQ ID NO:22:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 146 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide

FRIT SECTEMBE DESCRIPTION: SEC ID NO.23:

That His ier Ser Alz dan Hillys dan Ala Dal ler. Ny let Tro Ny I

typ Val Ash Val Asp 310 Val Bly 317 310 Ala Leo 317 Arg Leo Deu 20 \$25 \$30

Val Val Tyr Pro Trp Thr Gln Arg Phe Phe Glu Ser Phe Gly Asp Leu 35 40 45

for Ash Ale Ash Ala Val Met Gly Ash Pro Lys Val Lys Ala His Gly

Mal Leu Ala Arg Arg Leu Gly His Asp Phe Ash Pro Asp Mal Glm Ala 120 Ala Phe Gir Lys Val Val Ala Gly Val Ala Asn Ala Leu Ala His Lys 135 Tyr His 145 (2) INFORMATION FOR SEQ ID NO:23: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:23: Val His Let Ser Ala Glu Glu Lys Glu Ala Val Leu Gly Leu Trp Gly Lys Val Asn Val Asp Glu Val (2) INFORMATION FOR SEQ ID NO:24: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear \*.l+ ModECULE TYPE: peptide (x1) SECUENCE DESCRIPTION: SEC ID NO:24: Val Leu Ser Ala Ala Asp Lys Ala Asp Val Lys Ala Ala Trp Gly Lys 1 5 10 15 mai day day din er (IVER) de la comoció de la

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:25:

Phe Pro H.s Phe Asn Leu Ser His Gly Ser Asp Gln Val Lys
1 5 10

(2) INFORMATION FOR SEQ ID NO:26:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 10 amino acids
(B) TYPE: amino acid
(C) STRANDEDNESS:
(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:26:

Leu Val Val Tyr Pro Trp Thr Gln Arg Phe 1 5 10

(2) INFORMATION FOR SEQ ID NO:27:

(ii) MOLECULE TYPE: peptide

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 8 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- MIN SEQUENCE DESCRIPTION: SEA II NOIDE
- Tal Tal lyr Pro Irp Thr Win Ar c
- (E) INFORMATION FOR SEQ ID NO:28:
  - HIT SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 4 amino acids
    - (B) TYPE: amino acid

Ty: Pro Trp Thr

- (2) INFORMATION FOR SEQ ID NO:29:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 4 amino acids
    - (в) PYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide
  - (x1) SEQUENCE DESCRIPTION: SEQ ID NO:29:

Gly Tyr Pro Tyr

- (2) INFORMATION FOR SEQ ID NO:30:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 4 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:30:

Gly Lys Pro Tyr

THE IMPORMATION FOR SEQ ID NO.31:

- 1 OR MENUE OF ARACTERISTICS:
  - (A) LEWGTE: 4 amino acids
  - dry MYPE amino acid
  - (C) STRANDEDNESS:
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide

(C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:32: Lys Met Trp Met Phe Arg (2) INFORMATION FOR SEQ ID NO:33: (i) SEQUENCE CHAFACTERISTICS: (A) LENGTH: 8 amino acids (E) TYPE: amino acid (C) STRANDEDHESS: (D) TOPOLOGY: linear (ii) MODECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:33: Val Arg Arg Met Phe Gly Gly Tyr (2) INFORMATION FOR SEQ ID NO:34: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 6 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: single (10 TOPOLOGY: limear CHARTER INTERPOLER (xi) SEQUENCE DESCRIPTION: SEQ ID NO:34:

(B) TYPE: amino acid

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Phe Leu Gly Pne Pro Thr

THE PMATICAL FOR SECTION NOTES:

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(ix) FEATUPE:
                                (A) MAME KEY: Modified-site
                                (B) LOCATION. 1
                                (D) OTHER INFORMATION: product: "position 1"
          label- Xaa
         note: "pyroGln"
            (xi) SEQUENCE DESCRIPTION: SEQ ID NO:35:
               Maa Glr Glr Asp Cys Lys
1 5
(2) INFORMATION FOR SEQ ID NO:36:
                (i) SEQUENCE CHARACTERISTICS:
                                (A) LENGTH: 4 amino acids
                                 (B) TYPE: amino acid
                                 (C) STRANDEDNESS: single
                                 (D) TOPOLOGY: linear
            (ii) MODECULE TYPE: peptide
            (ix) FEATURE:
                               (A) NAME/KEY: Modified-site
                                 (B) LOCATION: 1
                                 (D) OTHER INFORMATION: /product= "position 1"
         label= Xaa
         note: "N-AbetylSer"
            (xi) SEQUENCE DESCRIPTION: SEQ ID NO:36:
                Xaa Asp Lys Pro
(2) INFORMATION FOR SEQ ID NO:37:
               HER SEQUEDCE CHARACTERISTICS:
                                (A) LENGTH: .3 amino acids
                                (E) TYPE: amino acid
                                (C) STRANDEDMESS: single
                                SIM TOPMIN'S Linear
             (ii) MOLECULE TYPE: peptide
                IN- FEATURE:
                               (A) NAME REY Modifieleste
                                GRO LOCATIONS 1
                                   TO MINISTER OF THE CONTROL OF THE STATE OF T
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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:37:

Xaa Pro His Xaa Asp Leu Ser His Gly Ser Ala Gln Val